



## INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

<b>(51) International Patent Classification <sup>7</sup> :</b> <b>A61K 45/06, A61P 35/00, A61K 41/00</b>	<b>A2</b>	<b>(11) International Publication Number:</b> <b>WO 00/37107</b> <b>(43) International Publication Date:</b> 29 June 2000 (29.06.00)
<b>(21) International Application Number:</b> PCT/US99/30776 <b>(22) International Filing Date:</b> 22 December 1999 (22.12.99) <b>(30) Priority Data:</b> 60/113,786                      23 December 1998 (23.12.98)      US <b>(71) Applicant (for all designated States except US):</b> G.D. SEARLE & CO. [US/US]; Corporate Patent Department, P.O. Box 5110, Chicago, IL 60680-5110 (US). <b>(72) Inventors; and</b> <b>(75) Inventors/Applicants (for US only):</b> MCKEARN, John, P. [US/US]; 18612 Bable Meadows Drive, Glencoe, MO 63038 (US). GORDON, Gary [US/US]; 3282 University Avenue, Highland, IL 60035 (US). CUNNINGHAM, James, J. [CA/US]; 3733 North Bell Avenue, Chicago, IL 60618 (US). GATELY, Stephen, T. [CA/US]; 357 E. Shady Pines Court, Palatine, IL 60067-8800 (US). KOKI, Alane, T. [US/US]; 6689 Highway 185, Beaufort, MO 63013 (US). MASFERRER, Jaime, L. [CL/US]; 1213 Blairshire, Ballwin, MO 63011 (US). <b>(74) Agents:</b> KEANE, J., Timothy et al.; G.D. Searle & Co., Corporate Patent Dept., P.O. Box 5110, Chicago, IL 60680-5110 (US).		<b>(81) Designated States:</b> AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).  <b>Published</b> <i>Without international search report and to be republished upon receipt of that report.</i>
<b>(54) Title:</b> METHOD OF USING A CYCLOOXYGENASE-2 INHIBITOR AND A MATRIX METALLOPROTEINASE INHIBITOR AS A COMBINATION THERAPY IN THE TREATMENT OF NEOPLASIA <b>(57) Abstract</b> <p>The present invention provides methods to treat or prevent neoplasia disorders in a mammal using a combination of a cyclooxygenase-2 inhibitor, a matrix metalloproteinase inhibitor and an antineoplastic agent.</p>		